Association between Oral Hygiene and Metabolic Syndrome

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Association between Oral Hygiene and Metabolic Syndrome: A Systematic Review and Meta-Analysis

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Abstract: Emerging evidence has linked poor oral hygiene to metabolic syndrome (MetS), but

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previously, no summary of evidence has been conducted on the topic. This systematic review and meta-analysis aims to evaluate the associations of oral hygiene status and care with MetS. A systematic search of the PubMed and Web of Science databases from inception to March 17, 2021, examination of reference lists was conducted to identify eligible observational studies. A random-effects model was applied to pool the effects of oral hygiene status and care on MetS. Thirteen studies met the inclusion criteria and had sufficient methodological quality. Good oral hygiene status (OR = 0.30 (0.13–0.66); I² = 91%), frequent tooth brushing (OR = 0.68 (0.58–0.80); I² = 89%), and frequent interdental cleaning (OR = 0.89 (0.81–0.99); I² = 27%) were associated with a lower risk of MetS. Only one study examined the association between dental visits and MetS (OR = 1.10 (0.77–1.55)). Our findings suggested that there might be inverse associations of oral hygiene status, tooth-brushing frequency, and interdental cleaning with MetS. However, substantial heterogeneity for tooth-brushing frequency and inconsistent results for oral hygiene status in subgroup analyses were observed. Ther

Keywords: oral hygiene; dental plaque; oral bacteria; tooth brushing; interdental cleaning; dental visit; metabolic syndrome

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1. Introduction

37 tabolic syndrome (MetS), a clustering of abdominal obesity, hyperglycemia, hypertension, and dyslipidemia, represents a growing public health concern globally [1]. Although the pressure of MetS differs depending on diagnostic criteria, age group, and ethnicity [58], it is estimated to affect around 25% of the world population [2,3]. MetS raises the risk of type 2 diabetes mellitus (T2DM) and cardiovascular diseases [1] and is associated with a 20% increase in healthcare costs [4].

63 Several risk factors for MetS have been identified. Besides socioeconomic status (SES) [5], smoking [6], diet [7], and physical activity [8], oral diseases, such as periodontal diseases and dental caries, are associated with MetS [9–11]. The link between oral and 61 temic diseases is suggested due to common risk factors, subgingival biofilm harboring Gran 62 gative bacteria, and periodontium serving as a cytokine reservoir [12].

Poor oral hygiene is the primary cause of common oral diseases. Accumulation of dental plaque allows bacterial growth that may lead to inflamed periodontal tissues and eventually create bacteremia and systemic inflammation [13,14]. Invading bacteria from severe caries or endodontic infections is also thought to provoke similar mechanisms [10,15,16]. Chronic low-grade inflammation underlies the development of metabolic

disorders [17,18], and a study found that systemic exposure to periodontal bacteria was associated with MetS [13].

Tooth brushing and interdental cleaning, which are the main forms of oral self-care, together with regular professional care, are important measures for plaque control or removal and maintaining optimal oral health [19–21]. Poor oral hygiene care is associated with low-grade inflammation [22], suggesting its potential link to MetS [23]. The association of poor oral hygiene care with a higher risk of the components of MetS, such as obesity [24], diabetes [25,26], hypertension [26,27], and dyslipidemia [26,28], as well as with cardiovascular disease [14,22], has been demonstrated.

Although several epidemiological studies have reported the association of oral hygiene 41 tus [29] and care [23,30] with MetS, some studies found no such association [31,32]. To date, there has not been a systematic review conducted on the topic. A summary of evidence can provide a better understanding of the potential relationship and help healthcare practitioners deliver more targeted care. It can provide more substance for the formulation of public health programs and policies, especially strategies for the prevention and management of MetS.

The aim of our study was to systematically review the association of oral hygiene status and care with MetS and to quantify the strength of associations.

2. Materials and Methods

The systematic review and meta-analysis were performed according to the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines [33]. The protocol was registered on the PROSPERO database (No. CRD42021243292) [34]. The research question was: Is better oral hygiene status or care associated with a lower risk of MetS?

2.1. Eligibility Criteria

The inclusion criteria were as follows: (1) The design of the study was cross-sectional, case—control, or cohort; (2) the exposure was oral hygiene status (e.g., oral hygiene index (OHI), plaque index (PI), plaque score (PSc)) or care (i.e., tooth brushing, interdental cleaning, and dental visit); (3 3 he outcome was MetS, clearly defined using diagnostic criteria for the condition (e.g., National Cholesterol Education Program Adult Treatment Panel III (NC 20 ATP III), International Diabetes Federation (IDF), Joint Interim Statement (JIS)); (4) the study assessed the assoc 20 ion between exposures and outcome in multiple analysis. There was no limitation on the characteristics of the study population. Animal studies, clinical trials, reviews, editorial letters, commentaries, case series, and case reports were excluded.

2.2. Search Strategy

A systematic search was performed on the PubMed and Web of Science databases, with the following keywords: oral hygiene, dental deposit, OHI, PI, PSc, tooth brushing, interdental cleaning, dental visit, and MetS. While no date restrictions were imposed, the language was limited to English. The last search was on March 17, 2021. Details of the search strategy can be seen in Supplementary Table S1. Examination of reference lists of eligible studies and relevant systematic reviews were also conducted to identify further relevant studies.

17 2.3. Study Selection and Data Extraction

Two authors independently screened all titles and abstracts to evaluate eligibility. Relevant studies were then examined for full-text review. Any ambiguities or disagreements were resolved by consensus. JabRef 5.2 was used during the review process.

Data from included studies were extracted independently at two authors using a data extraction form. The following information was collected: first author, publication year, study country, study design, sample size, age, gender, type of oral hygiene assessment, diagnostic criteria used for MetS, number of MetS cases, adjusted odds ratio (OR) or risk ratio (RR) with 95% confidence interval (CI), and adjustment factors. Discrepancies in data extraction were resolved by consensus.

2.4. Quality Assessment

Two authors independently examined the quality of included studies using the Newcastle–Ottawa Scale for cross-sectional, case–control, and cohort studies, as applicable. The three main domains examined were the selection of participants, comparability of study groups, and assessment of exposure/outcome of interest. The total scores for case–control and cohort studies were 9 points, while cross-captional studies were 8 points [35,36]. The included studies were then categorized into high (≥7 points), moderate (4–6 points), or low (0–3 points) quality. Any disagreements were resolved by consensus.

2.5. Statistical Analyses

Meta-analysis was conducted separately for different types of exposure (i.e., oral hygiene status, tooth brushing, and interdental cleaning). The OR was used as the common measure for the association between oral hygiene and MetS. The reported RR was considered approximately as OR [37]. The data utilized in the meta-analysis were the estimates and the corresponding 95% CI from the most adjusted model in the studies.

The categorization of exposure varied between studies. Poor oral hygiene status or care was used as the reference group, equivalent to the highest value of OHI, PI, and PSc or the lowest frequency category of tooth brushing, interdental cleaning, and dental visits in each study. If a study classified the exposure into more than two categories, a single effect estimate was produced by combining the results of the categories using a feffects (FE) model [38]. An overall pooled OR for the main analysis was calculated using a random-effects (RE) model (DerSimonian and Laird).

Heterogeneity was assessed using the F statistic, with the value of \geq 50% representing substantial heterogeneity [37,39]. Potential sources of heterogeneity were assessed using prespecified subgroup analyses by study design and country. E 67 lination of publication bias using funnel plot and Egger's test was only recommended if there were an adequate number of studies (>10) [40,41].

Meta-analysis was conducted using the generic inverse variance method in Review Manager (RevMan) 5.4 software (The Cochrane Collaboration, 2020) [42].



3.1. Literature Search

Figure 1 shows the process and the results of study selection. A total of 595 records were identified, of wh 24 144 were duplicates; 380 irrelevant studies were eliminated. Of the 71 22 dies selected for full-text review, 13 met the eligibility criteria and were included in the review and meta-analysis.

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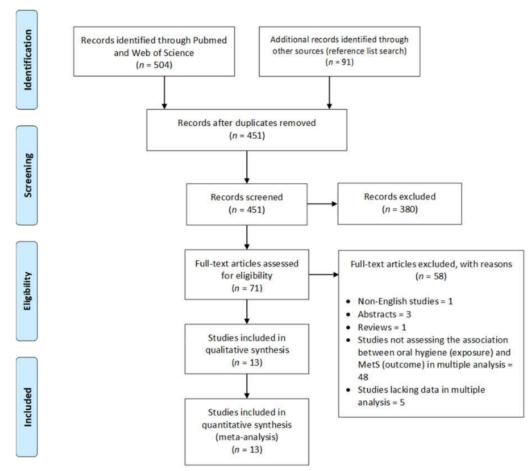


Figure 1. PRISMA flow diagram of the literature search and study selection [33]. MetS, metabolic syndrome.

3.2. Characteristics of Studies

Table 1 shows the main characteristics of the included studies. They consisted of seven cross-sectional, three case—control, and three cohort studies. A study by Shearer et al. [32] examined data from a cohort study. However, because our exposure of interest (modified OHI-S) was measured simultaneously with the outcome (MetS) at age 38, we chose to consider it as cross-sectional and reported the results of their cross-sectional model.

Eleven studies were from Asian countries, and one study each was from Finland and New Zealand. All were conducted among adult populations. Publication years ranged from 2009 to 2020, and the mean sample size was 4251.

Six studies reported oral hygiene status, six studies reported tooth-brushing frequency, two studies reported interdental cleaning, and one study reported dental visits as study factors. In the meta-analysis, a study by Tsutsumi et al. [43] was treated as two separate studies, as it reported the results independently for males and females instead of total samples. A similar approach was applied to a study by Kim et al. [44], as it provided separate data on interdental brushing and flossing.

Health examination was performed in all included studies to ascertain MetS conditions. Four studies used the NCEP ATP III criteria or its adapted version, five studies used JIS criteria, two used IDF criteria, and two used other criteria to define MetS. The most common confounders adjusted in the studies were age, gender, SES, smoking status, alcohol consumption, physical activity, and periodontal parameters. All studies reported a measure of associations as ORs, except for one study [31].

26 Table 1. Main characteristics of the 13 included studies.

Author,	j	Studen Designa	Sample Size	Age	Type of Oral	Diagnostic Criteria	Number	Statistical Analysis;	A 5000 to 1500 A
Fublication	Country	Study Design	(M, F)	Range	Hygiene	for MetS	of Cases	Adjustments	Association
Fukui et al., 2012 [45] Kim et al., 2013 [44]	Japan South Korea	Cross-sectional	6421 (M: 4944, F: 1477) 18742 (M: 8034, F: 10708)	34-77	Toothbrushing frequency (times/day) Toothbrushing frequency (frimes/day), 57 use of dental floss (yes or no), use of interdental brush (yes or no)	Modified NCEP ATP III*, except the use of BMI > 25 kg/m² to define obesity. Treatments for raised TG and reduced HDL were not recorded. Modified NCEP ATP III* for Asians.	958	Logistic regression; age, gender, smoking habit, alcohol consumption, C- reactive protein, number of teeth, periodontal parameter (PD or CAL). Logistic regression; age, gender, income, education, smoking, alcohol intake, and physical activities.	OR (95% CI) Adjusted by PD: 51 time daily (reference) 2 times daily = 0.67 (0.57–0.78) 23 times daily = 0.50 (0.40–0.64) Adjusted by CAL: 51 time daily [reference] 2 times daily = 0.50 (0.39–0.63) OR (95% CI) Tooth-brushing frequency: 23 times daily = 1.23 (1.04–1.47) 52 times daily = 1.23 (1.04–1.47) Use of dental floss: Yes [reference] No = 1.23 (1.07–1.41) Use of interdental brush: Yes [reference]

OR (95% CI) Males: None (reference) 1 time daily = 0.57 (0.40-0.81) 2 times daily = 0.50 (0.35-0.71) 23 times daily = 0.42 (0.29-0.61) Females: 1 time daily (reference) 2 times daily = 0.65 (0.48-0.87) 2 times daily = 0.44 (0.32-0.62)	OR (95% CI) Frequency of daily toothbrushing (continuous) = 0.887 (0.84-0.94)
Logistic regression; Males: age, exercise during holidays, favorite seasoning, eating soup, sugar in coffee, having an interest in losing weight, housekeeping during holidays; Females: age, favorite seasoning, worrying about job, sugar in coffee, pickles and food boiled in soy sauce, exercise during holidays, eating quickly, preparation of dinner, solving problems immediately.	Logistic regression; age, gender, household income, education, smoking, alcohol intake, physical activity, periodontitis.
3624	2834
Obesity (body mass percentage > 20% in men or > 30% in women, and/or BMI > 2.5 kg/m²) and at least one of the following: TG > 150 mg/dL and/or low mg/dL and/or low HDL < 40 mg/dL or drug for hypertriglyæridemia, SBP > 130 mm Hg and/or DBP ≥ 85 mm Hg or drug for hypertension, FPG > 110 mg/dL or drug for diabetes).	Three or more of the following five: WC ≥ 90 cm in men or ≥85 cm in women, TG > 150 mg/dL or treatment for raised TC, HDL < 40 mg/dL
Tooth- brushing frequency (times/day)	Tooth- brushing frequency (times/day)
30-59	35-79
12548 (M: 7703, F: 4845)	8314 (M: 3860, F: 4454)
Cross-sectional	Cross-sectional
Japan	South
Tsutsumi and Kakuma, 2015 [43]	Kim et al., 2019 [46]

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			OR (95% CI) None (reference) Sometimes = 1.19 (0.92–1.54) Everyday = 0.71 (0.55–0.92)	OR (95% CI) Low (reference) High = 0.95 (0.44, 2.01)
			Logistic regression; age, gender, smoking, exercise, weight gain, eating speed, cholesterol drug intake, community periodontal index, number of teeth.	Logistic regression; gender, low socioeconomic status, smoking,
			563	152
in men or \$0 mg/dL in women or	treatment for reduced HDL, SBP > 15 130 mm Hg and DBP ≥ 85 mm Hg or antihypertensive medication, FPG > 10	100 mg/dL or current use of antidiabetic medication.	JIS¹, except the use of BMI ≥ 25 kg/m² to define obesity and the use of HbA1c levels ≥ 5.6% to additionally define elevated glucose. Treatments for raised TG and reduced HDL were not included.	NCEP ATP III =, except the use of HbA1c ≥ 5.7% (≥39 mmol/mol) to define elevated glucose and
			Use of secondary oral hygiene products, such as dental floss or interdental brushes (none or sometimes or sometimes	Modified 32 OHI-S (very low (0-0.5) or low (>0.5-1.0) or moderate
			75 and 80	38
			2379 (M: 960, F: 1419)	836
			Cross-sectional	Cross-sectional
			Japan	New Zealand
			Saito et al., 2019 [47]	Shearer et al., 2018 [32]

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					(>1.0-1.5) or high (>1.5))	the use of antihypertensive drugs to additionally define elevated blood pressure.	dysglycemia, inflammatory load.	nia, tory load.	
Chen et al., 2011 [48]	Taiwan	Cross-sectional	253 (M:117, F: 136)	>18	Ы	Modified NCEP ATP III * for Asians, except the use of FPG ≥ 110 mg/dL or 145 previously diagnosed T2DM to define elevated glucose.		Logistic regression; age, gender, education, smoking, high-sensitivity C- reactive protein, and serum albumin.	OR (95% CI) PI score (continuous) = 1.724 (1.135–2.615)
Kobayashi et al., 2012 [30]	Japan	Cohort prospective, 3- year follow-up	685 (M: 513, F: 172)		Tooth- brushing frequency (times/day)	JIS [†] for Asians, except not induding treatments for raised 99 TG, reduced HDL, and elevated glucose.		Logistic regression; age, gender, smoking status, drinking status, breakfast eating, educational level, occupation (desk work), depressive symptoms, physical activity, and total caloric consumption.	OR (95% CI) \$\leq\$1 time daily (reference) 2 times daily = 0.80 (0.49-1.31) \$\leq\$3 times daily = 0.43 (0.19-0.97)
Tanaka et al., 2018 [23]	Japan	Cohort retrospective, 5- year follow-up	3722 (M: 2897, F: 825)	35-64	Tooth- brushing frequency (times/day),	JIS' for Asians, except the use of BMI ≥ 25 kg/m² to define obesity.	Logistic regression; age, gender, periodontal status, number of present	egression; er, al status, f present	OR (95% CI) Tooth-brushing frequency: ≤1 time daily (reference) 2 times daily = 0.83 (0.65–1.05)

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				dental check-			teeth, occupational	≥ 3 times daily = 0.64 (0.45–0.91)
				ups (regular			status, smoking	Dental check-ups:
				or irregular)			quantity, alcohol	Irregular (reference)
							consumption, physical	Regular = 1.10 (0.77-1.55)
							activity, dietary	
							behavior, food	
							preference, tooth-	
							brushing frequency,	
							dental check-ups, and	
							number of MetS	
							components at	
							baseline.	
							Poisson regression;	
							age, gender,	
							childhood BMI, family	
							income, adulthood	
		507 04.070 E.		Presence of			smoking (ever) and	RR (95% CI)
et al., Finland	prospective, 21-,	366 (M: 270, F:	27-43	visible plaque	JIS for Europeans.	153	socioeconomic status	No (reference)
2020 [31] 27-, 31-year	year	316)		(yes or no)			(education), and	Yes = $1.21 (0.87 - 1.86)$
dn-wonoi	dn						interaction terms	
							between caries and	
							periodontal	
							parameters.	
	-			PI (<2.5 or				OR (95% CI)
Pham, 2018 Vistoria (2006 – 2006	Ontrol	412 (M: 114, F:	70	2.51–2.90 or	TC # for A cione	200	Logistic regression;	<2.5 (reference)
	200,	298)	30-70	2.91–3.26 or	JO TOL ASIGILS.	200	age, gender.	2.51-2.90 = 4.81 (1.74-13.27)
control = 206)	= 206)			>3.27)				2.91–3.26 = 6.12 (2.24–16.70)

23.27 = 7.50 (2.80-20.12) OR (95% CI) ≤1 (reference) >1-1.5 = 4.81 (0.81-28.63) >1.5-2 = 13.06 (2.24-76.18) >2 = 47.4 (6.94-323.68)	OR (95% CI) PI score (continuous) = 14.69 (5.56–38.84)
Logistic regression; age, gender, smoking.	Logistic regression (backward); age, gender, smoking habits, bleeding index, PD, biomarkers (serum C-reactive protein, salivary IL-6,
152	114
IDF®	IDF §
PI (≤1 or >1-1.5 or >1.5-2 or >2)	Ы
37–78	37–78
208 (M: 85, F: 123)	163 (M: 60, F: 103)
Case—control (case = 152, control = 56)	Case–control (case = 114, control = 49)
China	China
Li et al., 2009 [49]	Li et al., 2020 [50]

M, male; F, female; MetS, metabolic syndrome; WC, 49 st circumference; BMI, body mass index; TG, triglycerides; HDL, high-density lipoprotein; SBP, systolist lood pressure; DBP, diastolic blood pressure; FPG, fasting plasma glucc 44 HbA1c, glycated haemoglobin; T2DM, type 2 diabets mellitus; OHI-S, simplified oral hygiene index; PI, plaque men or <50 mg/dL in women, blood pressure ≥ 130/85 mm Hg, FIT 110 mg/dL [51]. * The modified NCEP ATP III (2005) definition is a three of the following five: WC ≥ 102 cm (≥40 in) in men 8 ≥88 cm (≥35 in) in women (for Asigs: ≥90 cm (≥35 in) in men and ≥80 cm (≥31 in) in women), TC2 150 mg/dL (1.7 mmol/L) or treatment for reduced HDL, SBP ≥ 130 mm Hg or DBP ≥ 85 mm Hg or treatment for TC, HDL < 40 mg/dL (1.03 mmol/L) in men or <50 mg/dL (1.3 mmol/L) in women or treatment for reduced HDL, SBP ≥ 130 mm Hg or DBP ≥ 85 mm Hg or treatment for hypertension, FPG ≥ 100 mg/dL or treatment for 🚰 vated glucose [52]. § The International Diabetes Federational IDIA (2005) definition is increased WC (ethnicity specific) plus treatment for reduced HDL, SBP ≥ 130 mm Hg or DBP ≥ 85 mm Hg or treatment for hypertension, FPG ≥ 100 mg/dL (5.6 mmol/L) or previously diaphosed T2DM [53]. † The oint Interim Statement (JIS) (2009) definition is any three of the following five: increased WC (population- and country-specific), TG ≥ 150 mg/dL (1.7 mmol/L) or treatment index; PD, probing depth; CAL, clinical attachment level; OR, odds ratio; RR, risk ratio; CI, confidence interval. The National Cholesterol Education Program Adult Treatment Panel III (50)CEP ATP III) (2001) definition is any three of the following five: WC > 102 cm (>40 in) in men or >88 cm (>35 in) in women, TG ≥ 150 mg/dL, HDL < 40 mg/dL in any two of the following four: TG 2150 mg/dL (1.7 mmol/L) or treatment for raised TG, HDL < 40 mg/dL (1.03 mmol/L) in men or <50 mg/dL (1.29 mmol/L) in women or or raised TG, HDL < 40 mg/dL (1.0 mmol/L) in men or <50 mg/dL (1.3 mmol/L) in women or treatment for reduced HDL, SBP ≥ 130 mm Hg and/or DBP ≥ 85 mm Hg or treatment for hypertension, FPG ≥ 100 mg/dL or treatment for elevated glucose [54].

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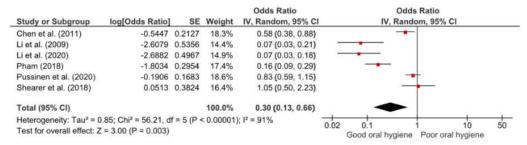
3.3. Quality Aspects of Studies

the included studies were of moderate to high quality. One cross-sectional study, two case-control studies, and three cohort studies were of high quality. Six cross-sectional studies and one case-control study were of moderate quality. Details of the quality assessment of included studies can be seen in Supplementary Table S2.

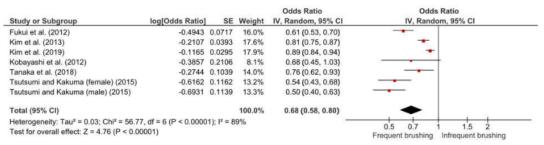
3.4. Association between Oral Hygiene Status, Care, and MetS

Figure 2 shows the results of the meta-analysis of associations of oral hygiene s_{38}^{-1} is, tooth-brushing frequency, and interdental cleaning with MetS. Good oral hygiene OR = 0.30; 95% CI = 0.13-0.66), frequent tooth brushing (OR = 0.68; 95% CI = 0.58-0.80), and frequent interdental cleaning (OR = 0.89; 95% CI = 0.81-0.99) were associated with a lower risk of MetS. While heterogeneity was minimal for interdental cleaning ($I^2 = 27\%$), there was substantial heterogeneity for oral hygiene status ($I^2 = 91\%$) and tooth-brushing frequency $I^2 = 89\%$).

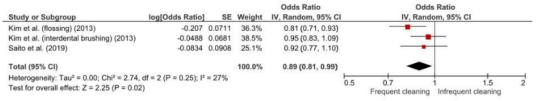
The association between dental visits a_{36} MetS was evaluated only in a study by Tanaka et al. It was found that dental visits were not significantly associated with MetS (OR = 1.10; 95% CI = 0.77–1.55) [23].



(a) Good versus poor oral hygiene status



(b) Frequent versus infrequent tooth brushing



(c) Frequent versus infrequent interdental cleaning

Figure 2. Meta-analysis of the associations of (a) oral hygiene status, (b) tooth-brushing frequency, and (c) interdental cleaning with metabolic syndrome.

3.5. Subgroup Analyses

Table 2 displays the results of subgroup analysis by study design for the association between oral hygiene status and MetS. The inverse association between oral hygiene status and MetS was only observed in the subgroup of case—control studies. Subgroup analysis by study design reduced heterogeneity to less than 50%.

Table 2. Subgroup analysis by study design for the association between oral hygiene status and MetS.

Subgrou	p	Number of Studies	OR (95% CI)	I ² (%)	p
Cross-section	onal	2	0.72 (0.41-1.26)	46	0.17
Case-cont	rol	3	0.11 (0.06-0.20)	39	0.19
Cohort		1	0.83 (0.59-1.15)	-	-

MetS, metabolic syndrome; OR, odds ratio; CI, confidence interval; I², percentage of variation due to heterogeneity; p, p-value for heterogeneity.

Table 3 shows the results of subgroup analyses for the association between tooth-brushing frequency and MetS. Frequent tooth brushing was consistently associated with a lower risk of MetS in all subgroup analyses. However, high heterogeneity was still observed among studies with a cross-sectional design. While subgroup analysis by country reduced heterogeneity, it remained above 50%.

Table 3. Subgroup analyses for the association between tooth-brushing frequency and MetS.

Subgroup	Number of Studies	OR (95% CI)	I ² (%)	р
Study design				
Cross-sectional	5	0.67 (0.55-0.81)	93	< 0.001
Cohort	2	0.74 (0.62-0.89)	0	0.64
Country				
Japan	5	0.61 (0.52-0.70)	55	0.06
g Korea	2	0.85 (0.78-0.93)	73	0.06

MetS, metabolic syndrome; OR, odds ratio; CI, confidence interval; I², percentage of variation due to heterogeneity; *p*, *p*-value for heterogeneity.

4. Discussion

Our systematic review and meta-analysis investigated as association of oral hygiene status and care with MetS. Better oral hygiene status, frequent tooth brushing, and frequent interdental cleaning were associated with a lower risk of MetS. However, substantial heterogeneity for tooth-brushing frequency and inconsistent results for oral hygiene status in subgroup analyses were noted. Our review identified only one study examining the association between dental visits and MetS, and found no association [23].

While our main analysis revealed an inverse association between better oral hygiene status and MetS, the finding was inconsistent in subgroup analysis by study design. Of all studies included in the meta-analysis for oral hygiene status, only studies by Shearer et al. [32] and Pussinen et al. [31], conducted in New Zealand and Finland, respectively, did not find an association. These different findings might be due to the age of the study samples. Both studies had relatively younger samples than the other studies, which had a sample mean age of more than 50 years. The stronger influence of periodontal inflammations on cardiometabolic health may only be observed in later life [32]. [31] reover, Pussinen et al. [31] reported both the adjusted RRs for MetS and β values for the number of MetS components. While the adjusted RR for [40] association between the presence of plaque and MetS was not significant, the β value for the association between the number of teeth with plaque and the number of MetS components was significant [31].

Our overall findings are in line with other systematic reviews and meta-analyses that demonstrated an association between oral health or hygiene and metabolic conditions [9,37]. Poor oral hygiene not only leads to dental infections, such as periodontitis, but it may also affect systemic health [55]. Periodontal bacteria in plaque, their products, and resulting local inflammatory response may enter the conditional description of the systemic inflammation [56]. Chronic exposure to proinflammatory cytokines, such as TNF- α may alter lipid metabolism, causing hyperlipidemia [57]. TNF- α may induce insulin resistance by directly affecting target organs (e.g., liver, muscle, and adipocytes) and by indirectly promoting the production of free fatty acids from adipocytes [58]. Elevated levels of proinflammatory cytokines may also contribute to pancreatic β -cells dysfunction, leading to the development of T2DM [57,59–61]. Moreover, recent evidence showed that *Porphyromonas gingivalis* might induce metabolic impairment by altering the gut microbiome [62].

Our study showed inverse relationships of tooth-brushing frequency and interdental cleaning with MetS. Despite substantial heterogeneity the findings of all subgroup analyses of tooth-brushing frequency were consistent. Tooth brushing is the most crucial self-care measure to control plaque and is a protective factor against periodontal diseases [63,64]. While a suggestion for proper frequency of tooth brushing could not be given, most of the included studies used a cut-off point of twice or more daily. Another review showed similar findings and indicated that brushing less than twice daily might not be beneficial for the prevention of DM [37]. In addition to tooth brushing, interdental cleaning is recommended for maintaining oral health. The daily use of interdental brushes was found to decrease periodontal bacteria, promote symbiotic microbiota, and reduce interdental inflammation [65]. It was suggested that poor oral hygiene could exaggerate Mets or increasing local and systemic inflammation [66].

An alternative explanation for the association between oral hygiene care and MetS might be that it is due to shared risk factors [14] or biased health consciousness. People with a healthier lifestyle might tend to adopt better oral hygiene care [67]. The fact that oral hygiene care may merely be an indicator of general health awareness or behaviors underscores the complexity of oral epidemiology [68]. However, most of the included studies in our review accounted for important confounders, such as age, gender, SES, smoking status, alcohol consumption, and physical activity, minimizing the bias.

The association between dental visits and MetS was not demonstrated in the study by Tanaka et al. [23]. This finding was similar to another study demonstrating no associations between dental visits, professional dental cleaning, and diabetes. It was argued that other confounders had more important roles in the development of diabetes than professional dental cleaning [25]. However, an earlier review has demonstrated the benefit of scaling and root planing on metabolic control and systemic inflammation reduction [23] patients with T2DM [69].

This systematic review and meta-analysis was the first to explore the association of oral hygiene status and care with MetS. The topic is seen as recent in the scientific literature, with the earliest identified studies published in 2009. It is also related to an emerging interest in the interrelationships between oral pathogens, oral microbiome dysbiosis, and systemic conditions [70]. Exploring this topic is relevant considering the importance of formulating policies with common risk factors approach to address both oral and general health [71]. Another strength of our review was the quality of the studies, which was moderate to high.

Our review might be limited by the methodological weakness of the included studies with a cross-sectional design. The number of cohort studies was also limited. Moreover, the restriction of studies to those published in English and the exclusion of a grey literature search might introduce bias. The risk of publication bias could not be ruled out and was not assessed in our study due to an inadeq 60: number of studies and high heterogeneity. Besides study design and country, the potential source of heterogeneity might be from the variability in measurement methods of oral hygiene status (e.g., the use

of different indices) and the reporting of tooth-brushing frequency and interdental cleaning between studies. Moreover, the criteria used to define MetS varied.

Information on tooth-brushing frequency and interdental cleaning was self-reported, which might be prone to bias. However, it might only be the type of nondifferential misclassification, leading to the underestimation of true effect estimates. Regular brushing does not necessarily reflect effective brushing, as the studies did not adjust for the duration and method of tooth brushing and the type of dentifrice used.

Finally, most of the included studies in our review were conducted among an Asian population, which may influence the generalizability of the findings worldwide. Further research conducted among other populations is warranted to provide more evidence. Using a uniform protocol for reporting oral hygiene (e.g., tooth-brushing frequency) may also facilitate better comparison.

5. Conclusions

Our study found that there might be inverse associations of oral hygiene status, tooth-brushing frequency, and interdental cleaning with MetS. However, substantial heterogeneity for tooth-brushing frequency and inconsistent results for oral hygiene status in subgroup analyses were observed. There was insufficient evidence on the association between dental visits and MetS. Further well-conducted studies, preferably of longitudinal design, are needed to confirm the associations of oral hygiene status and care with MetS and to explore their underlying mechanisms. Research on this topic will provide a valuable contribution to our current understanding of the interrelationship between oral health and MetS.

Supplementary Materials: The following are available online at www.mdpi.com/article/10.3390/jcm10132873/s1, Table S1: Database search strategy, Table S2: Quality assessment of the 13 included studies.

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References

- Cornier, M.A.; Dabelea, D.; Hernandez, T.L.; Lindstrom, R.C.; Steig, A.J.; Stob, N.R.; Van Pelt, R.E.; Wang, H.; Eckel, R.H. The metabolic syndrome. Endocr. Rev. 2008, 29, 777–822, doi:10.1210/er.2008-0024.
- Lear, S.A.; Gasevic, D. Ethnicity and metabolic syndrome: Implications for assessment, management and prevention. Nutrients 2020, 12, 15, doi:10.3390/nu12010015.
- Saklayen, M.G. The Global Epidemic of the Metabolic Syndrome. Curr. Hypertens. Rep. 2018, 20, doi:10.1007/s11906-018-0812-z.
- Curtis, L.H.; Hammill, B.G.; Bethel, M.A.; Anstrom, K.J.; Gottdiener, J.S.; Schulman, K.A. Costs of the metabolic syndrome in elderly individuals: Findings from the Cardiovascular Health Study. *Diabetes Care* 2007, 30, 2553–2558, doi:10.2337/dc07-0460.
- Blanquet, M.; Legrand, A.; Pélissier, A.; Mourgues, C. Socio-economics status and metabolic syndrome: A meta-analysis. Diabetes Metab. Syndr. Clin. Res. Rev. 2019, 13, 1805–1812, doi:10.1016/j.dsx.2019.04.003.

 Sun, K.; Liu, J.; Ning, G. Active Smoking and Risk of Metabolic Syndrome: A Meta-Analysis of Prospective Studies. PLoS ONE 2012, 7, doi:10.1371/journal.pone.0047791.

- Fabiani, R.; Naldini, G.; Chiavarini, M. Dietary patterns and metabolic syndrome in adult subjects: A systematic review and meta-analysis. Nutrients 2019, 11, 2056, doi:10.3390/nu11092056.
- 8. Joseph, M.S.; Tincopa, M.A.; Walden, P.; Jackson, E.; Conte, M.L.; Rubenfire, M. The impact of structured exercise programs on metabolic syndrome and its components: A systematic review. *Diabetes Metab. Syndr. Obes. Targets Ther.* **2019**, *12*, 2395–2404, doi:10.2147/DMSO.S211776.
- Gobin, R.; Tian, D.; Liu, Q.; Wang, J. Periodontal Diseases and the Risk of Metabolic Syndrome: An Updated Systematic Review and Meta-Analysis. Front. Endocrinol. 2020, 11, 1035–1057, doi:10.3389/fendo.2020.00336.
- Cao, X.; Wang, D.; Zhou, J.; Yuan, H.; Chen, Z. Relationship between dental caries and metabolic syndrome among 13 998 middle-aged urban Chinese. J. Diabetes 2017, 9, 378–385, doi:10.1111/1753-0407.12424.
- Ojima, M.; Amano, A.; Kurata, S. Relationship between decayed teeth and metabolic syndrome: Data from 4716 middle-aged male Japanese employees. J. Epidemiol. 2015, 25, 204–211, doi:10.2188/jea.JE20140132.
- Li, X.; Kolltveit, K.M.; Tronstad, L.; Olsen, I. Systemic diseases caused by oral infection. Clin. Microbiol. Rev. 2000, 13, 547–558, doi:10.1128/CMR.13.4.547-558.2000.
- Hyvärinen, K.; Salminen, A.; Salomaa, V.; Pussinen, P.J. Systemic exposure to a common periodontal pathogen and missing teeth are associated with metabolic syndrome. Acta Diabetol. 2015, 52, 179–182, doi:10.1007/s00592-014-0586-y.
- Chang, Y.; Woo, H.G.; Park, J.; Lee, J.S.; Song, T.J. Improved oral hygiene care is associated with decreased risk of occurrence for atrial fibrillation and heart failure: A nationwide population-based cohort study. Eur. J. Prev. Cardiol. 2020, 27, 1835–1845, doi:10.1177/2047487319886018.
- Gomes, M.S.; Blattner, T.C.; Sant'Ana Filho, M.; Grecca, F.S.; Hugo, F.N.; Fouad, A.F.; Reynolds, M.A. Can apical periodontitis modify systemic levels of inflammatory markers? A systematic review and meta-analysis. J. Endod. 2013, 39, 1205–1217, doi:10.1016/j.joen.2013.06.014.
- Scannapieco, F.A.; Cantos, A. Oral inflammation and infection, and chronic medical diseases: Implications for the elderly. Periodontol. 2000 2016, 72, 153–175, doi:10.1111/prd.12129.
- De Rooij, S.R.; Nijpels, G.; Nilsson, P.M.; Nolan, J.J.; Gabriel, R.; Bobbioni-Harsch, E.; Mingrone, G.; Dekker, J.M. Low-grade chronic inflammation in the relationship between insulin sensitivity and cardiovascular disease (RISC) population: Associations with insulin resistance and cardiometabolic risk profile. *Diabetes Care* 2009, 32, 1295–1301, doi:10.2337/dc08-1795.
- León-Pedroza, J.I.; González-Tapia, L.A.; del Olmo-Gil, E.; Castellanos-Rodríguez, D.; Escobedo, G.; González-Chávez, A. Low-grade systemic inflammation and the development of metabolic diseases: From the molecular evidence to the clinical practice. Cir. Cir. 2015, 83, 543–551, doi:10.1016/j.circen.2015.11.008.
- Claydon, N.C. Current concepts in toothbrushing and interdental cleaning. Periodontol. 2000 2008, 48, 10–22, doi: 10.1111/j.1600-0757.2008.00273.x.
- 20. Ainamo, J. Prevention of periodontal disease in the dental office. Int. Dent. J. 1984, 34, 56-61.
- Lim, L.P.; Davies, W.I.R. Comparison of various modalities of "simple" periodontal therapy on oral cleanliness and bleeding. J. Clin. Periodontol. 1996, 23, 595–600, doi:10.1111/j.1600-051X.1996.tb01830.x.
- De Oliveira, C.; Watt, R.; Hamer, M. Toothbrushing, inflammation, and risk of cardiovascular disease: Results from Scottish Health Survey. BMJ 2010, 340, 1400, doi:10.1136/bmj.c2451.
- Tanaka, A.; Takeuchi, K.; Furuta, M.; Takeshita, T.; Suma, S.; Shinagawa, T.; Shimazaki, Y.; Yamashita, Y. Relationship of toothbrushing to metabolic syndrome in middle-aged adults. J. Clin. Periodontol. 2018, 45, 538–547, doi:10.1111/jcpe.12876.
- Nijakowski, K.; Lehmann, A.; Rutkowski, R.; Korybalska, K.; Witowski, J.; Surdacka, A. Poor oral hygiene and high levels of inflammatory cytokines in saliva predict the risk of overweight and obesity. *Int. J. Environ. Res. Public Health* 2020, 17, 1–10, doi:10.3390/ijerph17176310.
- Chang, Y.; Lee, J.S.; Lee, K.J.; Woo, H.G.; Song, T.J. Improved oral hygiene is associated with decreased risk of new-onset diabetes: A nationwide population-based cohort study. *Diabetologia* 2020, 63, 924–933, doi:10.1007/s00125-020-05112-9.
- Fujita, M.; Ueno, K.; Hata, A. Lower frequency of daily teeth brushing is related to high prevalence of cardiovascular risk factors. Exp. Biol. Med. 2009, 234, 387–394, doi:10.3181/0809-RM-265.
- Choi, H.M.; Han, K.; Park, Y.-G.; Park, J.-B. Associations Among Oral Hygiene Behavior and Hypertension Prevalence and Control: The 2008 to 2010 Korea National Health and Nutrition Examination Survey. J. Periodontol. 2015, 86, 866–873, doi:10.1902/jop.2015.150025.
- Song, T.J.; Kim, J.W.; Kim, J. Oral health and changes in lipid profile: A nationwide cohort study. J. Clin. Periodontol. 2020, 47, 1437–1445, doi:10.1111/jcpe.13373.
- Pham, T. The association between periodontal disease severity and metabolic syndrome in Vietnamese patients. Int. J. Dent. Hyg. 2018, 16, 484–491, doi:10.1111/idh.12350.
- Kobayashi, Y.; Niu, K.; Guan, L.; Momma, H.; Guo, H.; Cui, Y.; Nagatomi, R. Oral health behavior and metabolic syndrome and its components in adults. J. Dent. Res. 2012, 91, 479

 –484, doi:10.1177/0022034512440707.
- Pussinen, P.J.; Paju, S.; Viikari, J.; Salminen, A.; Taittonen, L.; Laitinen, T.; Burgner, D.; Kahonen, M.; Lehtimaki, T.; Hutri-Kahonen, N.; et al. Childhood Oral Infections Associate with Adulthood Metabolic Syndrome: A Longitudinal Cohort Study. J. Dent. Res. 2020, 99, 1165–1173, doi:10.1177/0022034520929271.

32. Shearer, D.M.; Thomson, W.M.; Cameron, C.M.; Ramrakha, S.; Wilson, G.; Wong, T.Y.; Williams, M.J.A.; McLean, R.; Theodore, R.; Poulton, R. Periodontitis and multiple markers of cardiometabolic risk in the fourth decade: A cohort study. *Community Dent. Oral Epidemiol.* 2018, 46, 615–623, doi:10.1111/cdoe.12414.

- Moher, D.; Liberati, A.; Tetzlaff, J.; Altman, D.G. Preferred Reporting Items for Systematic Reviews and Meta-Analyses: The PRISMA Statement. PLoS Med. 2009, 6, e1000097, doi:10.1371/journal.pmed.1000097.
- 34. Santoso, C.M.A.; Ketti, F.; Nagy, A. Association between Oral Hygiene and Metabolic Syndrome: A Systematic Review and Meta-analysis. PROSPERO 2021 CRD42021243292. Available online: https://www.crd.york.ac.uk/prospero/display_record.php?ID=CRD42021243292 (accessed on 17 April 2021).
- Wells, G.; Shea, B.; O'Connell, D.; Peterson, J.; Welch, V.; Losos, M.; Tugwell, P. The Newcastle-Ottawa Scale (NOS) for Assessing the Quality of Nonrandomised Studies in Meta-Analyses. Available online: http://www.ohri.ca/programs/clinical_epidemiology/oxford.asp (accessed on 5 February 2021).
- Yuan, T.; Zou, H.; Zhao, J.; Yang, Z.; Li, L.; Cai, W.; Gu, J.; Hao, C.; Li, J.; Hao, Y.; et al. Circumcision to prevent HIV and other sexually transmitted infections in men who have sex with men: A systematic review and meta-analysis of global data. Artic. Lancet Glob. Health 2019, 7, e436–e447, doi:10.1016/S2214-109X(18)30567-9.
- 37. Fu, W.; Lv, C.; Zou, L.; Song, F.; Zeng, X.; Wang, C.; Yan, S.; Gan, Y.; Chen, F.; Lu, Z.; et al. Meta-analysis on the association between the frequency of tooth brushing and diabetes mellitus risk. Diabetes Metab. Res. Rev. 2019, 35, doi:10.1002/dmrr.3141.
- Bae, J.M. Comparison of methods of extracting information for meta-analysis of observational studies in nutritional epidemiology. Epidemiol. Health 2016, 38, e2016003, doi:10.4178/epih/e2016003.
- Analysing data and undertaking meta-analyses. In Cochrane Handbook for Systematic Reviews of Interventions Version 5.1.0 [updated March 2011]; Higgins, J.P., Green, S., Eds.; The Cochrane Collaboration: 2011.
- Addressing reporting biases. In Cochrane Handbook for Systematic Reviews of Interventions Version 5.1.0 [updated March 2011];
 Higgins, J.P.T., Green, S., Eds.; The Cochrane Collaboration: 2011.
- Alzahrani, H.; Mackey, M.; Stamatakis, E.; Zadro, J.R.; Shirley, D. The association between physical activity and low back pain: A systematic review and meta-analysis of observational studies. Sci. Rep. 2019, 9, 1–10, doi:10.1038/s41598-019-44664-8.
- Review Manager (RevMan) [Computer Program], Version 5.4; The Cochrane Collaboration: 2020. Available online: https://training.cochrane.org/online-learning/core-software-cochrane-reviews/revman/revman-non-cochrane-reviews (accessed on 16 March 2021).
- 43. Tsutsumi, C.; Kakuma, T. Regular Tooth Brushing is Associated with a Decreased Risk of Metabolic Syndrome According to a Medical Check-Up Database. *Kurume Med. J.* 2015, 61, 43–52, doi:10.2739/kurumemedj.MS64004.
- Kim, Y.-H.; Kim, D.-H.; Lim, K.S.; Ko, B.-J.; Han, B.-D.; Nam, G.-E.; Park, Y.-G.; Han, K.D.; Kim, J.-H.; Cho, K.-H. Oral health behaviors and metabolic syndrome: The 2008-2010 Korean National Health and Nutrition Examination Survey. Clin. Oral Investig. 2014, 18, 1517–1524, doi:10.1007/s00784-013-1112-2.
- Fukui, N.; Shimazaki, Y.; Shinagawa, T.; Yamashita, Y. Periodontal Status and Metabolic Syndrome in Middle-Aged Japanese. J. Periodontol. 2012, 83, 1363–1371, doi:10.1902/jop.2012.110605.
- Kim, J.S.; Kim, S.Y.; Byon, M.J.; Lee, J.H.; Jeong, S.H.; Kim, J.B. Association between periodontitis and metabolic syndrome in a korean nationally representative sample of adults aged 35–79 years. Int. J. Environ. Res. Public Health 2019, 16, 2930, doi:10.3390/ijerph16162930.
- Saito, M.; Shimazaki, Y.; Nonoyama, T.; Tadokoro, Y. Number of teeth, oral self-care, eating speed, and metabolic syndrome in an aged Japanese population. J. Epidemiol. 2019, 29, 26–32, doi:10.2188/jea.JE20170210.
- 48. Chen, L.-P.; Hsu, S.-P.; Peng, Y.-S.; Chiang, C.-K.; Hung, K.-Y. Periodontal disease is associated with metabolic syndrome in hemodialysis patients. *Nephrol. Dial. Transplant.* **2011**, *26*, 4068–4073, doi:10.1093/ndt/gfr209.
- Li, P.; He, L.; Sha, Y.Q.; Luan, Q.X. Relationship of Metabolic Syndrome to Chronic Periodontitis. J. Periodontol. 2009, 80, 541–549, doi:10.1902/jop.2009.080387.
- Li, P.; He, L.; Chen, Z.B.; Luan, Q.X. Biomarkers in Metabolic Syndrome Patients with Chronic Periodontitis. Chin. J. Dent. Res. 2020, 23, 191–197, doi:10.3290/j.cjdr.a45223.
- National Cholestrol Education Program (NCEP) Expert Panel on Detection Evaluation and Treatment of High Blood Cholestrol
 in Adults Treatment Panel III Third Report of the National Cholesterol Education Program (NCEP) Expert Panel on Detection,
 Evaluation, and Treatment of High Blood Cholesterol in Adults (Adult Treatment Panel III) final report. Circulation 2002, 106,
 3143–3421, doi:10.1161/circ.106.25.3143.
- Grundy, S.M.; Cleeman, J.I.; Daniels, S.R.; Donato, K.A.; Eckel, R.H.; Franklin, B.A.; Gordon, D.J.; Krauss, R.M.; Savage, P.J.; Smith, S.C.; et al. Diagnosis and management of the metabolic syndrome: An American Heart Association/National Heart, Lung, and Blood Institute scientific statement. Circulation 2005, 112, 2735–2752, doi:10.1161/CIRCULATIONAHA.105.169404.
- Alberti, K.G.M.M.; Zimmet, P.; Shaw, J. The metabolic syndrome A new worldwide definition. Lancet 2005, 366, 1059–1062, doi:10.1016/S0140-6736(05)67402-8.
- 54. Alberti, K.G.M.M.; Eckel, R.H.; Grundy, S.M.; Zimmet, P.Z.; Cleeman, J.I.; Donato, K.A.; Fruchart, J.C.; James, W.P.T.; Loria, C.M.; Smith, S.C. Harmonizing the Metabolic Syndrome: A Joint Interim Statement of The International Diabetes Federation Task Force on Epidemiology and Prevention; National Heart, Lung, and Blood Institute; American Heart Association; World Heart Federation; International Atherosclerosis Society; and International Association for the Study of Obesity. Circulation 2009, 120, 1640–1645, doi:10.1161/CIRCULATIONAHA.109.192644.

 Bui, F.Q.; Almeida-da-Silva, C.L.C.; Huynh, B.; Trinh, A.; Liu, J.; Woodward, J.; Asadi, H.; Ojcius, D.M. Association between periodontal pathogens and systemic disease. *Biomed. J.* 2019, 42, 27–35, doi:10.1016/j.bj.2018.12.001.

- Leite, F.R.M.; Nascimento, G.G. The Relationship Between Periodontal Diseases and Chronic Diseases. In Oral Epidemiology A
 Textbook on Oral Health Conditions, Research Topics and Methods; Peres, M.A., Antunes, J.L.F., Watt, R.G., Eds.; Springer: Cham,
 Switzerland, 2021; pp. 379–393.
- Iacopino, A.M. Periodontitis and diabetes interrelationships: Role of inflammation. Ann. Periodontol. 2001, 6, 125–137, doi:10.1902/annals.2001.6.1.125.
- Nishimura, F.; Iwamoto, Y.; Mineshiba, J.; Shimizu, A.; Soga, Y.; Murayama, Y. Periodontal Disease and Diabetes Mellitus: The Role of Tumor Necrosis Factor-α in a 2-Way Relationship. J. Periodontol. 2003, 74, 97–102, doi:10.1902/jop.2003.74.1.97.
- Wang, C.; Guan, Y.; Yang, J. Cytokines in the progression of pancreatic β-cell dysfunction. Int. J. Endocrinol. 2010, 2010, doi:10.1155/2010/515136.
- Cieślak, M.; Wojtczak, A.; Cieślak, M. Role of pro-inflammatory cytokines of pancreatic islets and prospects of elaboration of new methods for the diabetes treatment. Acta Biochim. Pol. 2015, 62, 15–21, doi:10.18388/abp.2014 853.
- Grover, H.S.; Luthra, S. Molecular mechanisms involved in the bidirectional relationship between diabetes mellitus and periodontal disease. J. Indian Soc. Periodontol. 2013, 17, 292–301, doi:10.4103/0972-124X.115642.
- Watanabe, K.; Katagiri, S.; Takahashi, H.; Sasaki, N.; Maekawa, S.; Komazaki, R.; Hatasa, M.; Kitajima, Y.; Maruyama, Y.; Shiba, T.; et al. *Porphyromonas gingivalis* impairs glucose uptake in skeletal muscle associated with altering gut microbiota. *FASEB J.* 2020, 35, e21171, doi:10.1096/fj.202001158R.
- Lertpimonchai, A.; Rattanasiri, S.; Arj-Ong Vallibhakara, S.; Attia, J.; Thakkinstian, A. The association between oral hygiene and periodontitis: A systematic review and meta-analysis. Int. Dent. J. 2017, 67, 332–343, doi:10.1111/idj.12317.
- Zimmermann, H.; Zimmermann, N.; Hagenfeld, D.; Veile, A.; Kim, T.S.; Becher, H. Is frequency of tooth brushing a risk factor for periodontitis? A systematic review and meta-analysis. Community Dent. Oral Epidemiol. 2015, 43, 116–127, doi:10.1111/cdoe.12126.
- Bourgeois, D.; Bravo, M.; Llodra, J.C.; Inquimbert, C.; Viennot, S.; Dussart, C.; Carrouel, F. Calibrated interdental brushing for the prevention of periodontal pathogens infection in young adults—A randomized controlled clinical trial. Sci. Rep. 2019, 9, doi:10.1038/s41598-019-51938-8.
- 66. Kim, S.W.; Cho, K.H.; Han, K.D.; Roh, Y.K.; Song, I.S.; Kim, Y.H. Tooth loss and metabolic syndrome in South Korea: The 2012 Korean national health and nutrition examination survey. *Medicine* 2016, 95, doi:10.1097/MD.000000000003331.
- 67. Yeung, C.A. Gums and heart disease: Healthy gums, healthy heart? BMJ 2010, 341, 113, doi:10.1136/bmj.c3710.
- Franchini, R.; Petri, A.; Migliario, M.; Rimondini, L. Poor oral hygiene and gingivitis are associated with obesity and overweight status in paediatric subjects. J. Clin. Periodontol. 2011, 38, 1021–1028, doi:10.1111/j.1600-051X.2011.01770.x.
- Baeza, M.; Morales, A.; Cisterna, C.; Cavalla, F.; Jara, G.; Isamitt, Y.; Pino, P.; Gamonal, J. Effect of periodontal treatment in patients with periodontitis and diabetes: Systematic review and meta-analysis. J. Appl. Oral Sci. 2020, 28, doi:10.1590/1678-7757-2019-0248.
- Santarelli, A.; Wong, D.T.W.; Lo Muzio, L. Editorial: Saliva and Oral Microbiota: From Physiology to Diagnostic and Therapeutic Implications. Front. Physiol. 2021, 11, 637599, doi:10.3389/fphys.2020.637599.
- Souza, M.L.; Massignan, C.; Peres, K.G.; Peres, M.A. Association between metabolic syndrome and tooth loss: A systematic review and meta-analysis. J. Am. Dent. Assoc. 2019, 150, 1027–1039.e7, doi:10.1016/j.adaj.2019.07.023.

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